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| TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371 | | ATTORNEY'S DOCKET NUMBER 020884-000009 |
| INTERNATIONAL APPLICATION NO. PCT/EP2004/000339 | | U.S. APPLICATION NO. (if known, see 37 CFR 1.5) |
| INTERNATIONAL FILING DATE January 19, 2004 | PRIORITY DATE CLAIMED January 19, 2004 | |
| TITLE OF INVENTION METHOD FOR OBTAINING A SINGULAR CELL MODEL CAPABLE OF REPRODUCING IN VITRO THE METABOLIC IDIOSYNCRASY OF HUMANS | | |
| APPLICANT(S) FOR DO/EO/US José Vicente Castell Ripoll María José Gómez-Lechón Ramiro Jover Atienza Agustín Lahoz Rodríguez | | |
| Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: | | |
| <ol style="list-style-type: none"> 1. <input checked="" type="checkbox"/> This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. 2. <input type="checkbox"/> This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. 3. <input checked="" type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below. 4. <input checked="" type="checkbox"/> The US has been elected (Article 31). 5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371(c)(2)) <ol style="list-style-type: none"> a. <input type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau). b. <input checked="" type="checkbox"/> has been communicated by the International Bureau. c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office(RO/US) 6. <input type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). <ol style="list-style-type: none"> a. <input type="checkbox"/> is attached hereto. b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4). 7. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)) <ol style="list-style-type: none"> a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau). b. <input type="checkbox"/> have been communicated by the International Bureau. c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired. d. <input checked="" type="checkbox"/> have not been made and will not be made. 8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). 9. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). 10. <input type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)). | | |
| Items 11 to 20 below concern document(s) or information included: | | |
| <ol style="list-style-type: none"> 11. <input type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98. 12. <input type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 13. <input checked="" type="checkbox"/> A preliminary amendment. 14. <input type="checkbox"/> An Application Data Sheet under 37 CFR 1.76. 15. <input type="checkbox"/> A substitute specification. 16. <input type="checkbox"/> A power of attorney and/or change of address letter. 17. <input type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825. 18. <input checked="" type="checkbox"/> A second copy of the published International Application under 35 U.S.C. 154(d)(4). 19. <input type="checkbox"/> A second copy of the English language translation of the International application under 35 U.S.C. 154(d)(4). | | |

This collection of information is required by 37 CFR 1.414 and 1.491-1.492. The information is required to obtain or retain a benefit by the public, which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and 1.14. This collection is estimated to take 15 minutes to complete, including gathering information, preparing, and submitting the completed form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Mail Stop PCT, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.

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| U.S. APPLICATION NO. (if known, see 37 CFR 1.5) | | INTERNATIONAL APPLICATION NO. PCT/EP2004/000339 | | ATTORNEY'S DOCKET NUMBER 020684-000009 | |
| 20. Other items or information: Copy of International Preliminary Examination Report; Copy of Reply to Written Opinion and Article 34 Amendment to the claims | | | | | |
| The following fees have been submitted | | | | CALCULATIONS PTO USE ONLY | |
| 21. <input checked="" type="checkbox"/> Basic national fee (37 CFR 1.492(a)).....\$300 | | | | \$ 300 | |
| 22. <input checked="" type="checkbox"/> Examination fee (37 CFR 1.492(c)) | | | | \$ 200 | |
| If the written opinion prepared by ISA/US or the International preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4).....\$0 | | | | | |
| All other situations.....\$200 | | | | | |
| 23. <input checked="" type="checkbox"/> Search fee (37 CFR 1.492(b)) | | | | \$ 400 | |
| If the written opinion prepared by ISA/US or the International preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4).....\$0 | | | | | |
| Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority.....\$100 | | | | | |
| International Search Report prepared by an ISA other than the US and provided to the Office or previously communicated to the US by the IB.....\$400 | | | | | |
| All other situations.....\$500 | | | | | |
| TOTAL OF 21, 22 and 23 = | | | | \$ 900 | |
| <input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing in compliance with 37 CFR 1.82(c) or (e) or computer program listing in an electronic medium) (37 CFR 1.492(i)). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. | | | | | |
| Total Sheets | Extra sheets | Number of each additional 50 or fraction thereof (round up to a whole number) | RATE | | |
| - 100 = | /50 = | | X \$250 | \$ | |
| Surcharge of \$130.00 for furnishing any of the search fee, examination fee, or the oath or declaration after the date of commencement of the national stage (37 CFR 1.492(h)). | | | | \$ | |
| CLAIMS | NUMBER FILED | NUMBER EXTRA | RATE | \$ | |
| Total Claims | 12 - 20 = | 0 | x \$50 | \$ 0 | |
| Independent Claims | 2 - 3 = | 0 | x \$200 | \$ 0 | |
| MULTIPLE DEPENDENT CLAIM(S) (if applicable) | | | + \$360 | \$ | |
| TOTAL OF ABOVE CALCULATIONS = | | | | \$ 900 | |
| <input checked="" type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2. | | | | | |
| SUBTOTAL = | | | | \$ 450 | |
| Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(i)). | | | | \$ | |
| TOTAL NATIONAL FEE = | | | | \$ 450 | |
| Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.26, 3.31). \$40.00 per property | | | | \$ | |
| TOTAL FEES ENCLOSED = | | | | \$ 450.00 | |
| | | | | Amount to be refunded: | \$ |
| | | | | Amount to be charged: | \$ |

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- a. ☐ A check in the amount of \$ _____ to cover the above fees is enclosed.
- b. ☐ Please charge my Deposit Account No. 13-4365 in the amount of \$ _____ to cover the above fees.
A duplicate copy of this sheet is enclosed.
- c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 13-4365. A duplicate copy of this sheet is enclosed.
- d. ☒ Fees are to be paid by Electronic Funds Transfer.

NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the International Application to pending status.

SEND ALL CORRESPONDENCE TO:

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27709
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Tristan A. Fuierer
SIGNATURE

Tristan A. Fuierer
NAME

52,926
REGISTRATION NUMBER

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

| | | | |
|--|---|----------------------|-------------------------|
| In re United States Patent Application of: |) | Docket No.: | 020884-9 |
| Applicant(s): |) | Examiner: | Not Yet Assigned |
| Application No.: |) | Art Unit: | Not Yet Assigned |
| New U.S. National Stage Application of PCT International Application No. PCT/EP2004/000339 |) | | |
| Int'l Filing Date: |) | Customer No.: | |
| 19 January 2004 |) | | |
| Priority Date: |) | | |
| 19 January 2004 |) | | |
| Title: |) | 24239 | |
| METHOD FOR OBTAINING A SINGULAR CELL MODEL CAPABLE OF REPRODUCING IN VITRO THE METABOLIC IDIOSYNCRASY OF HUMANS |) | | |

**SUBMISSION UNDER 35 U.S.C. §371 OF UNITED STATES PATENT APPLICATION
(NATIONAL PHASE PROCEEDINGS) BASED ON INTERNATIONAL APPLICATION NO.
PCT/EP2004/000339**

Mail Stop PCT
Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Sir:

Submitted herewith for filing under the provisions of 37 CFR §1.53 and 35 U.S.C. §371 is the above-referenced patent application, based on International Patent Application No. PCT/EP2004/000339. A copy of the PCT International Application, Response to Written Opinion and Article 34 Amendment to the Claims, the International Preliminary Report on Patentability, a Preliminary Amendment, and 35 U.S.C. §371 transmittal forms are included herewith.

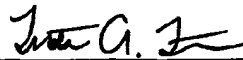
Please direct correspondence relating to this application to Tristan Fuierer, Moore & Van Allen, PLLC, P.O. Box 13706, Research Triangle Park, NC 27709, and direct telephonic communications relating to this application to Tristan Fuierer at (919) 286-8090.

Respectfully submitted,

MOORE & VAN ALLEN PLLC

Date: July 19, 2006

By:



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